

Patent Department 2292 Faraday Avenue Carlsbad, CA 92008 Facsimile Information

Sender's Name: Pamela D. Grooms

Isis Ref. No.: ISIS-3455 Date January 7, 2004 Serial Number: 09/295,463 Filing Date: April 13, 1999

Art Unit: 1631

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This Fax is from

Name: Pamela Grooms

Fax # (760) 603-3820 Telephone # (760) 603-2415

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PAGE 1/19* RCVD AT 1/7/2004 7:38:45 PM [Eastern Standard Time] * SVR:USPTO-EFXRF-1/2* DNIS:8729306 * CSID:760 603 3820 * DURATION (mm-ss):04-10

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	Application Number	09/295,463
TRANSMITTAL	Filing Date	April 13, 1999
FORM	First Named Inventor	Lex M. Cowsert
(to be used for all correspondence after initial	riling) Art Unit	1631
	Examiner Name	Ardin H. Marschel
Total Number of Pages In This Submission	Attorney Docket Number	I\$IS-345S
	ENCLOSURES (Check all th	at apply)
Fee Transmittal Form Fee Attached Amendment/Reply After Final Affidavits/declaration(s) Extension of Time Request Express Abandonment Request Information Disclosure Statement Certified Copy of Priority Document(s) Response to Missing Parts/ Incomplete Application Response to Missing Parts under 37 CFR 1.52 or 1,53	Drawing(s) Licensing-related Papers Petition Petition to Convert to a Provisional Application Power of Attorney, Revocation Change of Correspondence Add Terminal Disclaimer Request for Refund CD, Number of CD(s) Remarks Revocation of Power of Attorney with S	Other Enclosure(\$) (please Identify below):
SIGNA	TURE OF APPLICANT, ATTORN	IEY, OR AGENT
Firm or Individual name Signature Date	1/3/04	
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I hereby certify that this correspondence is be sufficient postage as first class mail in an em- the date shown below.	eling facsimile transmitted to the USPTO o	r deposited with the United States Postal Service with stents, P.O. Box 1450, Alexandria, VA 22313-1450 on
Typed or printed name Pamela Grooms		
Signature Demillo	-Lume	Date 17704

This collection of information is required by 37 CFR 1.5. The information is required to obtain or retain a benefit by the public which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.14. This collection is estimated to 12 minutes to complete, including gathering, preparing, and submitting the completed application form to the USPTO. Time will vary depending upon the individual case, Any comments on the amount of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, U.S. Department of Commerce, P.O. Box 1450, Alexandria, VA 22313-1450. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.

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REVOCATION OF POWER OF ATTORNEY WITH NEW POWER OF ATTORNEY AND CHANGE OF CORRESPONDENCE ADDRESS

sportd to a collection of information unles	as it displays a valid QMB control number
Application Number	09/295,463
Filing Date	April 13, 1999
First Named Inventor	Lex M. Cowsert
Art Unit	1631
Examiner Name	Ardin H. Marschel
Attorney Docket Number	ISIS-3455

I hereby revoke all previo	ous powers of attorney given in	ı the above-ider	itified application.
A Power of Attorney	is submitted herewith.		
OR I hereby appoint the	practitioners associated with the	Customer Numb	er: 35807
Please change the correspondence address for the above-identified application to: The address associated with			
Customer Number	er:	35807	
Firm or Individual Name			
Address			
Address			
City		State	Zip
Country	,		
Telephone		Fax	
I am the: Applicant/Inventor. Assignee of record of the entire interest. See 37 CFR 3.71. Statement under 37 CFR 3.73(b) is enclosed. (Form PTO/SB/96)			
	SIGNATURE of Applicant of	or Assignee of R	ecord
Name B. Lynne Parshall			
Signature			
Date 1 - 5	- 04	Telephone	760 603-2460
NOTE: Signatures of all the inventors or assignees of record of the entire interest or their representative(s) are required. Submit multiple forms if more than one signal pre is required, see below.			
Total of 2 forms	are submitted.		

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STATEMENT UNDER 37 CFR 3,73(b) Isis Pharmaceuticals In Applicant/Patent Owner: _ Application No./Patent No.: DA |Filed/Issue Date: _ Entitled: Identification of Genetic Targets-fix Modulation by Oligonulleothdes and generation of disprudent des for some Modulation USis Pharmaceuticals, Inc (Type of Assignee, e.g., corporation, partnership, university, government agency, etc.) (Name of Assignee) states_that it is: 1. It the assignee of the entire right, title, and interest; or 2. \square an assignee of less than the entire right, title and interest. The extent (by percentage) of its ownership interest is in the patent application/patent identified above by virtue of either: A. Di An assignment from the inventor(s) of the patent application/patent identified above. The assignment was recorded in the United States Patent and Trademark Office at Reel <u>O10047</u>, Frame <u>0285</u>, or for which a copy thereof is attached. ORB. [] A chain of title from the inventor(s), of the patent application/patent identified above, to the current assignee as shown below: 1. From: -The document was recorded in the United States Patent and Trademark Office at _, or for which a copy thereof is attached. ______, Frame Reel 2. From: To: The document was recorded in the United States Patent and Trademark Office at , Frame __, or for which a copy thereof is attached. Reel 3. From: The document was recorded in the United States Patent and Trademark Office at or for which a copy thereof is attached. [] Additional documents in the chain of title are listed on a supplemental sheet. [] Copies of assignments or other documents in the chain of title are attached. [NOTE: A separate copy (i.e., the original assignment document or a true copy of the original document) must be submitted to Assignment Division in accordance with 37 CFR Part 3, if the assignment is to be recorded in the records of the USPTO. See MPEP 302.08] The undersigned (whose title is supplied below) is authorized to act on behalf of the assignee. -5-04 Typed or printed name 760 603 - 24 Signature Telephone number

This collection of information is required by 37 CFR 3.73(b). The information is required to obtain or retain a benefit by the public which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.14. This collection is estimated to take 12 minutes to complete, including gathering, preparing, and submitting the completed application form to the USPTO. Time will vary depending upon the individual case. Any comments on the amount of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, U.S. Department of Commerce, P.O. Box 1450, Alexandria, VA 22313-1450. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450,

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AUGUST 26, 199 Mackiewicz & North LLP

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PAUL K. LEGAARD ONE LIBERTY PLACE - 46TH FLOOR

1650 MARKET STREET

PHILADELPHIA, PENNSYLVANIA 19103-7301



1515.3455

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RECORDATION DATE: 06/24/1999

REEL/FRAME: 010047/0285

NUMBER OF PAGES: 13

BRIEF: ASSIGNMENT OF ASSIGNOR''S INTEREST (SEE DOCUMENT FOR DETAILS).

ASSIGNOR:

COWSERT, LEX M.

DOC DATE: 05/26/1999

ASSIGNOR:

BAKER, BRENDA F.

DOC DATE: 06/15/1999

ASSIGNOR:

MCNEIL, JOHN

DOC DATE: 06/02/1999

ASSIGNOR:

FREIER, SUSAN M.

DOC DATE: 06/09/1999

ASSIGNOR:

SASMOR, HENRI M.

DOC DATE: 06/11/1999

ASSIGNOR:

BROOKS, DOUGLAS G.

DOC DATE: 06/03/1999

ASSIGNOR:

OHASHI, CARA

· ...

DOC DATE: 05/24/1999

010047/0285 PAGE 2

ASSIGNOR:

WYATT, JACQUELINE R.

DOC DATE: 05/26/1999

ASSIGNOR:

BORCHERS, ALEXANDER H.

DOC DATE: 06/01/1999

ASSIGNOR:

VICKERS, TIMOTHY A.

DOC DATE: 05/27/1999

ASSIGNEE:

ISIS PHARMACEUTICALS, INC. 2292 FARADAY AVENUE CARLSBAD, CALIFORNIA 92008

SERIAL NUMBER: 09295463

PATENT NUMBER:

FILING DATE: 04/13/1999

ISSUE DATE:

KIMBERLY WHITE, EXAMINER ASSIGNMENT DIVISION OFFICE OF PUBLIC RECORDS

06-28-1999



RECORDATION FORM COVER SHEET PATENTS ONLY

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101077363

Patents and Trademarks. Please record the attached original documents or copy i

1. Name of conveying party(ies):

Lex M. Cowsert, Brenda F. Baker, John McNeil, Susan M. Freier, Henri M. Sasmor, Douglas G. Brooks, Cara Ohashi, Jacqueline R. Wyatt, Alexander H. Borchers, Timothy A. Vickers

Additional name(s) of conveying party(ies) attached? () YES (X) NO

- 3. Nature of Conveyance:
 - (X) Assignment
- () Merger
- () Security Agreement
- () Change of Name
- () Other:

Execution Date(s): May 26, 1999; June 15, 1999; June 2, 1999; June 9, 1999; June 11, 1999; June 3, 1999; May 24, 1999; May 26, 1999; June 1, 1999; May 27, 1999

2. Name and address of receiving party(les):

Name: ISIS Pharmaceuticals, Inc.

Street Address: 2292 Faraday Avenue

City: Carlsbad

State California

ZIP 92008

Additional name(s) & address attached? ()YES (X)NO

- 4. Application number(s) or patent number(s):09/295,463
 If this document is being filed together with a new application, the execution date of the application is:
 - A. Patent Application No.(s)

B. Patent No.(s)

Additional numbers attached? () YES (X) NO

5. Name and address of party to whom correspondence concerning documents should be mailed:

Name: Paul K. Legaard

Internal Address: WOODCOCK WASHBURN KURTZ

MACKIEWICZ & NORRIS LLP One Liberty Place - 46th Floor

Philadelphia, Pennsylvania 19103-7301

Street Address: WOODCOCK WASHBURN KURTZ

MACKIEWICZ & NORRIS LLP One Liberty Place - 46th Floor

1650 Market Street

Philadelphia, Pennsylvania 19103;7301

- 6. Total number of applications and patents involved: One
- 7. Total fee (37 CFR 3.41):.....\$40.00 (X) Enclosed
 - () Authorized to be charged to Deposit Account Number 23-3050
- 8. Please charge any deficiency or credit any overpayment to Deposit Account Number 23-3050

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9. Statement and signature.

To the best of my knowledge and belief, the foregoing information is true and correct and any attached copy is a true copy of the original document.

Paul K. Legaard/Reg. 38,534

Name of Person Signing/Reg. No.

Park Face Signature

June 21, 1999

Date

Total number of pages including cover sheet, attachments, and document: 14

Mail documents to be recorded with required cover sheet information to:

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Washington, D.C. 20231

PATENT

ASSIGNMENT

WHEREAS, Lex M. Cowsert of Carlsbad, California, Brenda F. Baker of Carlsbad, California, John McNeil of LaJolla, California, Susan M. Freier of San Diego, California, Ocean Side 115 Henri M. Sasmor of Encinitas, California, Douglas G. Brooks, of San Marcos, California, Cara Ohashi of San Francisco, California, Jacqueline R. Wyatt of Encinitas, California, Alexander H. Borchers of Encinitas, California and Timothy A. Vickers of Oceanside, California, have invented a certain invention entitled IDENTIFICATION OF GENETIC TARGETS FOR MODULATION BY OLIGONUCLEOTIDES AND GENERATION OF OLIGONUCLEOTIDES FOR GENE MODULATION, for which we have made application for Letters Patent of the United States, said application having been filed on April 13, 1999 and assigned Serial No. 09/295,463; and

WHEREAS, ISIS Pharmaceuticals, Inc. hereinafter referred to as the assignee, of 2292 Faraday Avenue, Carlsbad CA, a corporation of Delaware, is desirous of acquiring the entire right, title and interest in and to the said inventions or improvements and in and to the said application, and in, to and under any and all Letters Patent which may be granted on or as a result thereof in any and all countries:

NOW, THEREFORE, for and in consideration of the sum of One Dollar (\$1.00) to each of us in hand paid by said assignee, and other good and valuable consideration, the receipt of which is hereby acknowledged, we, the said assignors, have sold, assigned, transferred and set over, and by these presents do hereby sell, assign, transfer and set over to said assignee, the entire right, title and interest in and to said inventions or improvements and said application and any and all continuations, divisions and renewals of and substitutes for said application, and in, to and under any and all Letters Patent which may be granted on or as a result thereof in the United States and any and all other countries, and any reissue or reissues or extension or extensions of said Letters Patent, and assign

ASSIGNMENT - PAGE 1 OF 12

PATENT

to and authorize said assignce, to file in our names applications for Letters Patent in all countries, the same to be held and enjoyed by said assignee, its successors, assigns, nominees or legal representatives, to the full end of the term or terms for which said Letters Patent respectively may be granted, reissued or extended, as fully and entirely as the same would have been held and enjoyed by us had this assignment, sale and transfer not been made.

I

AND we hereby covenant that we have full right to convey the entire interest herein assigned. and that we have not executed and will not execute any agreement in conflict herewith, and we further covenant and agree that we will each time request is made and without undue delay, execute and deliver all such papers as may be necessary or desirable to perfect the title to said inventions or improvements, said application and said Letters Patent to said assignee, its successors, assigns, nominees, or legal representatives, and each of us agrees to communicate to said assignee or to its nominee all known facts respecting said inventions or improvements, said application and said Letters Patent, to testify in any legal proceedings, to sign all lawful papers to execute all disclaimers and divisional, continuing, reissue and foreign applications, to make all rightful oaths, and generally to do everything possible to aid said assignee, its successors, assigns, nominees and legal representatives to obtain and enforce for its or their own benefit proper patent protection for said inventions or improvements in any and all countries.

AND we hereby authorize and request the Commissioner of Patents and Trademarks of the United States and any official of any country or countries foreign to the United States whose duty it is to issue patents on applications as aforesaid, to issue to said assignee, as assignee of the entire right, title and interest, any and all Letters Patent for said inventions or improvements, including any and all Letters Patent of the United States which may be issued and granted on or as a result of the application aforesaid, in accordance with the terms of this assignment.

ASSIGNMENT - PAGE 2 OF 12

PATENT

IN WITNESS WHEREOF, we have hereunto set our lands and seals.

State of California County of San Diego

day of __, 1999, before me, the undersigned, a Notary Public for the state, personally appeared (Lex M. Cowsert) personally known to me/provedto-me on the basis of satisfactory evidence to be the person who subscribed the foregoing Assignment and acknowledged to me that he she executed the same in his her authorized capacity, and that by his her signature on the Assignment the person, or the entity on behalf of which the person acted, executed the Assignment.

Signature of Notary

LANE B.D. SCHWARTZ Commission # 1196197 Notary Public - California San Diego County My Comm. Expires Sep 13, 2002

WITNESS my hand and official seal.

ASSIGNMENT - PAGE 3 OF 12

PATENT

Dated:

6/15/99

Brenda F. Baker

State of California

County of Can Diego

County of San Diego

SS.



WITNESS my hand and official seal.

Ι

DOCKET NO.: ISIS-3455	P.	ATENT
Dated: 6/2/99, 1999	John McNeil	
State of California)	•	
County of San Diego) ss.		
On this day of day of Notary Public for the state, personally appeared (a me on the basis of satisfactory evidence to be the pand acknowledged to me that he she executed the his her signature on the Assignment the person, of executed the Assignment.	same in his har muthorized	mment

LANE B.D. SCHWARTZ
Commission # 1196197
Notary Public - Collionia
San Diego County
My Comm. Expires Sep 13, 2002

WITNESS my hand and official scal.

Signature of Notary

Ø 013

DOCKET NO.: ISIS-3455

PATENT

Dated: 9-June 1999

Susan M. Freier

State of California

SS.

County of San Diego

LANE B.D. SCHWARTZ
Commission # 1196197
Notary Public - California
San Diego County
My Comm. Expires Sep 13, 2002

WITNESS my hand and official seal.

Signature of Notary

Ι

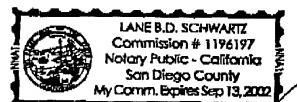
DOCKET NO.: ISIS-3455	PATENT
Dated: 6-11-99 1999	Hun Susner
He	enri M. Sasmor
State of California)	
County of San Diego) ss.	
- voury i abito for areatate, personally appeared;	Henri M. Sasmor) personally known to me/proved
to me on the basis of satisfactory evidence	to be the person who subscribed the foregoing
and that by his her signature on the Assignment	e executed the same in his/her authorized capacity, at the person, or the entity on behalf of which the
person acted, executed the Assignment.	A my 12 man of which the

LANE B.D. SCHWARTZ Commission # 1196197 Notary Public - California San Diego Caunty Ny Comm. Expires Sep 13, 2002 WITNESS my band and official seal.

ASSIGNMENT - PAGE 7 OF 12

DOCKET NO.: ISIS-3455		PATENT
Dated:6/3/99	, 1999 Douglas G. Brooks	- 122.02(12
State of California		
County of San Diego) ss.)	
On this 3rd	day of, 1999, before me, the under	ersigned, a

Signature of Notary



WITNESS my hand and official seal.

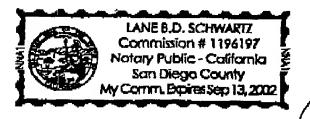
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Signature of Notary

SUNITA RAO
Comm. # 1175447
NOTARY PUBLIC - CALIFORNIA
San Mateo County
My Comm. Expires March 6, 2002

DOCKET NO.: ISIS-3455			PATENT
Dated:5/24	, 1999	Jacqueline R. Wyatt	_
State of California)		
County of San Diego) ss.)		

Signature of Notary



WITNESS my hand and official seal.

ASSIGNMENT - PAGE 10 OF 12

PATENT

Dated: June

State of California

Sŝ.

County of San Diego

_____, 1999, before me, the undersigned, a day of_ Notary Public for the state, personally appeared (Alexander H. Borchers) personally known to me/proved to me on the basis of satisfactory evidence to be the person who subscribed the foregoing Assignment and acknowledged to me that he she executed the same inches her authorized capacity, and that by his/her signature on the Assignment the person, or the entity on behalf of which the person acted, executed the Assignment.

Signature of Notary

LANE B.D. SCHWARTZ Commission # 1196197 iotary Public - California san Diego County My Comm. Brokes Sep 13, 2002 WITNESS my hand and official seal.

ASSIGNMENT - PAGE 11 OF 12

County of San Diego

DOCKET NO.: ISIS-3455

Dated: 5(27), 1999

Timothy A. Vickers

State of California)

SS.

On this day of May 1999, before me, the undersigned, a Notary Public for the state, personally appeared (Timothy A. Vickers) personally known to me/proved to me on the basis of satisfactory evidence to be the person who subscribed the foregoing Assignment and acknowledged to me that he she executed the same in hig/her authorized capacity, and that by hig/her signature on the Assignment the person, or the entity on behalf of which the person acted, executed the Assignment.

WITNESS my hand and official seal.

LANE B.D. SCHWARTZ
Commission # 1196197
Notary Public - Collifornia
San Diega County
My Comm. Expires Sep 13, 2002

Signature of Notary